REPORT DOCUMENTATION PAGE

Form Approved

OMB No. 0704-0188

Public reporting burden for this collection of information is estimated to average 1 hour per response, including the time for reviewing instructions, searching existing data soul gathering and maintaining the data needed, and completing and reviewing the collection of information. Send comments regarding this burden estimate or any other aspect of collection of information, including suggestions for reducing this burden, to Washington Headquarters Services, Directorate for information Operations and Reports, 1215 Jeffe Davis Highway, Suite 1204, Arlington, VA 22202-4302, and to the Office of Management and Budget, Paperwork Reduction Project '9704-0188), Washington, DC 20503.

1. AGENCY USE ONLY (Leave blank)

2. REPORT DATE May 1990 3. REPORT TYPE AND DATES COVERED

Final

15 Mar 1987 - 24 Mar 1990

A TITLE AND SUBTITLE

Development of Microencapsulation Techniques

5. FUNDING NUMBERS

DAAL03-87-K-0044

6. AUTHOR(S)

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PERFORMING ORGANIZATION REPORT NUMBER

9. SPONSORING / MONITORING AGENCY NAME(S) AND ADDRESS(ES)

7. PERFORMING ORGANIZATION NAME(S) AND ADDRESS(ES)

California Institute of Technology

U. S. Army Research Office

P. O. Box 12211

Research Triangle Park, NC 27709-2211

10. SPONSORING / MONITORING AGENCY REPORT NUMBER

ARO 23967.5-CH

11. SUPPLEMENTARY NOTES

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12b. DISTRIBUTION CODE

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13. ABSTRACT (Maximum 200 words)

Surfactant vesicles are small, spherical shell-like structures composed of bilayers of surfactant molecules which can be used to contain an aqueous solution. Since a variety of materials can be encapsulated in such vesicles, they are ideal vehicles for many different applications. The advantages of vesicle formulations involve protection from the environment, sustained release, and targeted delivery of vesicle contents. Such potential for phospholipid vesicles as chemical delivery systems, and possibly many other roles has motivated a significant effort towards

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14. SUBJECT TERMS

15. NUMBER OF PAGES

Microencapsulation Techniques, Membrane Stability, Vesicle Fusion, Permeability, Lipid Analogues, Disulfide Polymerization, 16. PRICE COOE

Cholesterol Derivatives, Chr. 194 E. M. 1. SECURITY CLASSIFICATION 18. SECURITY CLASSIFICATION SECURITY CLASSIFICATION OF REPORT

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SECURITY CLASSIFICATION OF ABSTRACT

20. LIMITATION OF ABSTRAC

NSN 7540-01-280-5500

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Standard Form 298 (Rev. 2-89) Prescribed by ANSI Std. 239-18 298-102

improving their properties. The primary objectives for much of the current research in this area have therefore been to obtain enhanced membrane stability (mechanical and chemical) and decreased or controlled vesicle fusion and permeability. Major progress toward these ends has been made by the introduction of polymerizeable lipid analogues and to a lesser extent with the formation of polymer coated vesicles.

Studies have been made of the effects of disulfide polymerization and the incorporation of cholesterol derivatives on the structure and properties (permeability, stability, and size) of phospholipid vesicles. Studies have been initiated on the interaction of polymers with liposomes.

FINAL TECHNICAL REPORT

John D. Baldeschwieler Professor of Chemistry

14 May 1990

U. S. ARMY RESEARCH OFFICE

For

Contract DAAL03-87-K-0044

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Final Report

DEVELOPMENT OF MICROENCAPSULATION TECHNIQUES

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Surfactant vesicles are small, spherical shell-like structures composed of bilayers of surfactant molecules which can be used to contain an aqueous solution. Since a variety of materials can be encapsulated in such vesicles, they are ideal vehicles for many different applications. The advantages of vesicle formulations involve protection from the environment, sustained release, and targeted delivery of vesicle contents. Such potential for phospholipid vesicles as chemical delivery systems, and possibly many other roles has motivated a significant effort towards improving their properties. The primary objectives for much of the current research in this area have therefore been to obtain enhanced membrane stability (mechanical and chemical) and decreased or controlled vesicle fusion and permeability. Major progress toward these ends has been made by the introduction of polymerizeable lipid analogues(1) and to a lesser extent with the formation of polymer coated vesicles(2).

With support of the Army Research Office (Grant No. DAAL03-87-K-0044) we have explored the effects of disulfide polymerization and incorporation of cholesterol derivatives on the structure and properties (permeability, stability, and size) of phospholipid vesicles. We have also initiated studies of the interaction of polymers with liposomes.

The detailed results of these studies have been published or submitted for publication as follows:

- 1. Handel, Tracy M., "Disulfide Polymerizeable Phosphatidylcholines: Characterization of Membrane Physical Properties and Investigations of *in vivo* Behavior". Ph.D. diss., California Institute of Technology, Pasadena, CA, 1989.
- 2. Goodrich, Raymond P., "Membrane Bound Carbohydrates: An Approach for Stabilization During Freezing and Drying". Ph.D. Diss., California Institute of Technology, Pasadena, CA. 1990.
- 3. Goodrich, Raymond P., Handel, Tracy M. <u>BBA</u> <u>938</u>, 143-154, 1988.
- 4. Goodrich, Raymond P., Crowe, John H., Crowe, Lois M., Baldeschwieler, John D. <u>Biochem</u>. 1990, submitted. "Alterations in Membrane Surfaces Induced by Attachment of Carbohydrates".

- Goodrich, Raymond P., John D. Baldeschwieler, <u>BBA</u> 1990, submitted.
 "Protection of Vesicles Against Damage During Freeze Drying by Addition of Membrane Associated Carbohydrate Derivatives".
- 6. Goodrich, Raymond P., Baldeschwieler, John D., <u>Cryobiology</u> 1990, submitted. "The Cryoprotective Action of Synthetic Glycolipids".

I have attached copies of the abstracts of the two theses (nos. 1. and 2.) as well as preprints of the three manuscripts which have recently been submitted for publication.

List of scientific personnel:

Chris DiSimone Mitsuko Fujiwara Raymond P. Goodrich, Ph.D., 1990 Tracy M. Handel, Ph.D., 1989 Wilton Vannier, M.D., Ph.D.



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